Application No.:

NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 C.F.R. 1.821 - 1.825 for the following reason(s):

owing reason(s).	
1. This application clearly fails to comply with the requirements of 37 C.F.R. 1.821-1.825. Applicant's attention is directed to these regulations, published at 1114 OG 29, May 15, 1990 and at 55 FR 18230, May 1, 1990.	>
2. This application does not contain, as a separate part of the disclosure on paper copy, a "Sequence Listing" as required by 37 C.F.R. 1.821(c).	е
3. A copy of the "Sequence Listing" in computer readable form has not been submitted as required by 37 C.F.R. 1.821(e).	y
4. A copy of the "Sequence Listing" in computer readable form has been submitted. However, the content of the computer readable form does not comply with the requirements of 37 C.F.R. 1.822 and/or 1.823, as indicated on the attached copy of the marked-up "Raw Sequence Listing."	<u>.</u>
5. The computer readable form that has been filed with this application has been found to be damaged and/or unreadable as indicated on the attached CRF Diskette Problem Report. A Substitute computer readable form must be submitted as required by 37 C.F.R. 1.825(d).	je
6. The paper copy of the "Sequence Listing" is not the same as the computer readable form of the "Sequence Listing" as required by 37 C.F.R. 1.821(e).	
7. Other:	
Applicant Must Provide: An initial or substitute computer readable form (CRF) copy of the "Sequence Listing".	
An initial or substitute paper copy of the "Sequence Listing", as well as an amendment directing its entry into the specification.	
A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 C.F.R. 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d).	
For questions regarding compliance to these requirements, please contact	
For Rules Interpretation, call (703) 308-4216	

For CRF Submission Help, call (703) 308-4212

For Patentin software help, call (703) 308-6856

PLEASE RETURN A COPY OF THIS NOTICE WITH YOUR RESPONSE

OIPE

```
RAW SEQUENCE LISTING
                                       DATE: 07/30/2001
PATENT APPLICATION: US/09/462,416
                                      TIME: 14:57:53
Input Set : A:\PTO_VSK.txt
```

Output Set: N:\CRF3\07302001\I462416.raw

```
ENTERED
      3 <110> APPLICANT: REVEL, Michel
             CHEBATH, Judith
             LAPIDOT, Tsvee
             KOLLET, Orit
      8 <120> TITLE OF INVENTION: CHIMERIC INTERLEUKIN-6 SOLUBLE RECEPTOR/LIGAND PROTEIN,
ANALOGS THEREOF
             AND USES THEREOF
    11 <130> FILE REFERENCE: REVEL=15
     13 <140> CURRENT APPLICATION NUMBER: 09/462,416
    14 <141> CURRENT FILING DATE: 2000-04-13
    16 <150> PRIOR APPLICATION NUMBER: PCT/IL98/00321
    17 <151> PRIOR FILING DATE: 1998-07-09
     19 <150> PRIOR APPLICATION NUMBER: IL 121284
     20 <151> PRIOR FILING DATE: 1997-07-10
    22 <150> PRIOR APPLICATION NUMBER: IL 122818
    23 <151> PRIOR FILING DATE: 1997-12-30
     25 <160> NUMBER OF SEQ ID NOS: 12
     27 <170> SOFTWARE: PatentIn version 3.1
     29 <210> SEQ ID NO: 1
     30 <211> LENGTH: 13
     31 <212> TYPE: PRT
    32 <213> ORGANISM: Artificial Sequence
     34 <220> FEATURE:
     35 <223> OTHER INFORMATION: synthetic \_
     37 <400> SEQUENCE: 1
     39 Glu Phe Gly Ala Gly Leu Val Leu Gly Gly Gln Phe Met
     43 <210> SEQ ID NO: 2
    44 <211> LENGTH: 22
    45 <212> TYPE: DNA
    46 <213> ORGANISM: Artificial Sequence
    48 <220> FEATURE:
    49 <223> OTHER INFORMATION: synthetic
    51 <400> SEQUENCE: 2
                                                                               22
    52 ctagtgggcc cggggtggcg gg
    55 <210> SEQ ID NO: 3
    56 <211> LENGTH: 25
    57 <212> TYPE: DNA
    58 <213> ORGANISM: Artificial Sequence
    60 <220> FEATURE:
    61 <223> OTHER INFORMATION: synthetic
    63 <400> SEQUENCE: 3
                                                                               25
    64 gactagtagc tatgaactcc ttctc
    67 <210> SEQ ID NO: 4
    68 <211> LENGTH: 21
    69 <212> TYPE: DNA
    70 <213> ORGANISM: Artificial Sequence
```

72 <220> FEATURE:

RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/462,416

DATE: 07/30/2001 TIME: 14:57:53

21

31

Input Set : A:\PTO_VSK.txt

Output Set: N:\CRF3\07302001\I462416.raw

- 73 <223> OTHER INFORMATION: synthetic
- 75 <400> SEQUENCE: 4
- 76 agggccattt gccgaagagc c
- 79 <210> SEQ ID NO: 5
- 80 <211> LENGTH: 31
- 81 <212> TYPE: DNA
- 82 <213> ORGANISM: Artificial Sequence
- 84 <220> FEATURE:
- 85 <223> OTHER INFORMATION: synthetic
- 87 <400> SEQUENCE: 5
- 88 gatccgggcg ġcgggggagg ggggcccggg c
- 91 <210> SEQ ID NO: 6
- 92 <211> LENGTH: 14
- 93 <212> TYPE: PRT
- 94 <213> ORGANISM: Artificial Sequence
- 96 <220> FEATURE:
- 97 <223> OTHER INFORMATION: synthetic
- 99 <400> SEQUENCE: 6
- 101 Gly Gly Gly Gly Asp Pro Gly Gly Gly Gly Gly Pro Gly
 - 02 1 5 1
- 105 <210> SEQ ID NO: 7
- 106 <211> LENGTH: 543
- 107 <212> TYPE: PRT
- 108 <213> ORGANISM: Artificial Sequence
- 110 <220> FEATURE:
- 111 <223> OTHER INFORMATION: synthetic '
- 113 <400> SEQUENCE: 7
- 115 Met Leu Ala Val Gly Cys Ala Leu Leu Ala Ala Leu Leu Ala Ala Pro
 - 10 1
- 119 Gly Ala Ala Leu Ala Pro Arg Arg Cys Pro Ala Gln Glu Val Ala Arg
 - 20 20 25
- 123 Gly Val Leu Thr Ser Leu Pro Gly Asp Ser Val Thr Leu Thr Cys Pro
 - 24 35 40
- 127 Gly Val Glu Pro Glu Asp Asn Ala Thr Val His Trp Val Leu Arg Lys
- .28 50 55
- 131 Pro Ala Ala Gly Ser His Pro Ser Arg Trp Ala Gly Met Gly Arg Arg
- 132 65 70 75 80
- 135 Leu Leu Leu Arg Ser Val Gln Leu His Asp Ser Gly Asn Tyr Ser Cys
- 136 85 90 95
- 139 Tyr Arg Ala Gly Arg Pro Ala Gly Thr Val His Leu Leu Val Asp Val
- 140 100 105 110 143 Pro Pro Glu Glu Pro Gln Leu Ser Cys Phe Arg Lys Ser Pro Leu Ser
- 144 115 120 125
- 147 Asn Val Val Cys Glu Trp Gly Pro Arg Ser Thr Pro Ser Leu Thr Thr
- 148 130 135 140
- 151 Lys Ala Val Leu Leu Val Arg Lys Phe Gln Asn Ser Pro Ala Glu Asp
- 152 145 150 155 . 160
- 155 Phe Gln Glu Pro Cys Gln Tyr Ser Gln Glu Ser Gln Lys Phe Ser Cys 156 165 170 175

RAW SEQUENCE LISTING DATE: 07/30/2001 PATENT APPLICATION: US/09/462,416 TIME: 14:57:53

Input Set : A:\PTO_VSK.txt

Output Set: N:\CRF3\07302001\1462416.raw

159 160	Gln	Leu	Ala	Val 180	Pro	Glu	Gly	Asp	Ser 185	Ser	Phe	Tyr	Ile	Val 190	Ser	Met
163 164		Val	Ala 195	Ser	Ser	Val	Gly	Ser 200	Lys	Phe	Ser	Lys	Thr 205	Gln	Thr	Phe
167 168	Gln	Gly 210	_	Gly			Gln 215	Pro	Asp	Pro	Pro	Ala 220	Asn	Ile	Thr	Val
	Thr 225	Ala	Val	Ala	Arg	Asn 230	Pro	Arg	Trp	Leu	Ser 235	Val	Thr	Trp	Gln	Asp 240
175 176	Pro	His	Ser	Trp	Asn 245	Ser	Ser	Phe	Tyr	Arg 250	Leu	Arg	Phe	Glu	Leu 255	Arg
179 180	Tyr	Arg	Ala	Glu 260	Arg	Ser	Lys		Phe 265	Thr	Thr	Trp	Met	Val 270	Lys	Asp
183 184	Leu	Gln	His 275	His	Cys	Val	Ile	His 280	Asp	Ala	Trp	Ser	Gly 285	Leu	Arg	His
187 188	Val	Val 290	Gln	Leu	Arg	Ala	Gln 295	Glu	Glu	Phe	Gly	Gln 300	Gly	Glu	Trp	Ser
	Glu 305	Trp	Ser	Pro	Glu	Ala 310	Met	Gly	Thr	Pro	Trp 315	Thr	Glu	Ser	Arg	Ser 320
195 196	Pro	Pro	Ala	Glu	Asn 325	Glu	Val	Ser	Thr	Pro 330	Met	Gln	Ala	Leu	Thr 335	Thr
199 200	Asn	Lys	Asp	Asp 340	Asp	Asn	Ile	Leu	Phe 345	Arg	Asp	Ser	Ala	Asn 350	Ala	Thr
203 204	Ser	Leu	Pro 355	Val	Glu	Phe	Met	Pro 360	Val	Pro	Pro	Gly	Glu 365	Asp	Ser	Lys
207 208	Asp	Val 370	Ala	Ala	Pro	His	Arg 375	Gln	Pro	Leu	Thr	Ser 380	Ser	Glu	Arg	Ile
	Asp 385	Lys	Gln	Ile	Arg	Tyr 390	Ile	Leu	Asp	Gly	Ile 395	Ser	Ala	Leu	Arg	Lys 400
215 216	Glu	Thr	Cys	Asn	Lys 405	Ser	Asn	Met	Cys	Glu 410	Ser	Ser	Lys	Glu	Ala 415	Leu
219 220	Ala	Glu	Asn	Asn 420	Leu	Asn	Leu	Pro	Lys 425	Met	Ala	Glu	Lys	Asp 430	Gly	Cys
223 224	Phe	Gln	Ser 435	_	Phe	Asn	Glu	Glu 440	Thr	Cys	Leu	Val	Lys 445		Ile	Thr
227 228	Gly	Leu 450	Leu	Glu	Phe	Glu	Val 455	Tyr	Leu	Glu	Tyr	Leu 460	Gln	Asn	Arg	Phe
	Glu 465	Ser	Ser	Glu	Glu	Gln 470	Ala	Arg	Ala	Val	Gln 475	Met	Ser	Thr	Lys	Val 480
235 236		Ile	Gln			Gln		Lys					Asp		Ile 495	
239 240	Thr	Pro	Asp	Pro 500	Thr	Thr	Asn	Ala	Ser 505	Leu	Leu	Thr	Lys	Leu 510	Gln	Ala
243 244	Gln	Asn	Gln 515	Trp	Leu	Gln	Asp	Met 520	Thr	Thr	His	Leu	Ile 525	Leu	Arg	Ser
247 248	Phe	Lys 530	Glu	Phe	Leu	Gln	Ser 535	Ser	Leu	Arg	Ala	Leu 540	Arġ	Gln	Met	
	51 <210> SEQ ID NO: 8															
	252 <211> LENGTH: 471 253 <212> TYPE: PRT															
233	~~	11		* ***			•									

RAW SEQUENCE LISTING DATE: 07/30/2001 PATENT APPLICATION: US/09/462,416 TIME: 14:57:53

Input Set : A:\PTO_VSK.txt
Output Set: N:\CRF3\07302001\1462416.raw

			RGAN:		Art	ific	ial :	Sequ	ence							
256	<220> FEATURE:															
257	<22	3> 0	THER	INF	ORMA!	rion	: sy	nthe	tic							
			EQUÉ													
261	Met	Asn	Ser	Phe	Ser	Thr	Ser	Ala	Phe	Gly	Pro	Val	Ala	Phe	Ser	Leu
262	1				5					10					15	
	Gly	Leu	Leu		Val	Leu	Pro	Ala		Phe	Pro	Ala	Pro		Pro	Pro
266	a 1	a 1	•	20	-		**- 7		25	D	TT! _		a 1	30	T	m l
269	_	Glu	Asp 35	ser	ьys	Asp	Val	A1a 40	Ala	Pro	HIS	Arg	45	Pro	Leu	Thr
273	Ser	Ser	Glu	Arg	Ile	Asp	Lys	Gln	Ile	Arg	Tyr	Ile	Leu	Asp	Gly	Ile
274		50					55					60				
277	Ser	Ala	Leu	Arg	Lys	Glu	Thr	Cys	Asn	Lys	Ser	Asn	Met	Cys	Glu	Ser
278	65					70					75					80
281	Ser	Lys	Glu	Ala	Leu	Ala	Glu	Asn	Asn	Leu	Asn	Leu	Pro	Lys	Met	Ala
282					85					90					95	
	Glu	Lys	Asp	Gly	Cys	Phe	Gln	Ser		Phe	Asn	Glu	Glu		Cys	Leu
286				100					105				•	110		
	Val	Lys		Ile	Thr	Gly	Leu		Glu	Phe	Glu	Val		Leu	Glu	Tyr
290			115					120				_	125			_
	Leu		Asn	Arg	Phe	Glu		Ser	Glu	Glu	Gln		Arg	Ala	Val	Gln
294		130	_		_		135					140				
		Ser	Thr	Lys	Val		Ile	Gln	Phe	Leu		Lys	Lys	Ala	Lys	
	145	_			_,	150	_	_	_	1	155	_		_	_	160
	Leu	Asp	Ala	He		Thr	Pro	Asp	Pro		Thr	Asn	Ala	Ser	Leu	Leu
302		_	_		165	~ 1	_	~ 1	_	170	-3	_		em 1	175	
	Thr	Lys	Leu		Ата	GIn	Asn	GIn		Leu	GIn	Asp	мет		Thr	HIS
306	T	T 1.	T	180	C	nh a	T	a 1	185	т	<i>c</i> 1-			190	3	71~
	Leu	тте		Arg	ser	Pne	ьуs		Pne	Leu	GIII	ser		ьeu	Arg	Ald
310	T 011	7 ~~	195	Mot	C1**	C1	C1	200	7 00	Dro	C1	C1	205	C1**	C1	cirr
314	ьeu	210	GIII	мес	GIY	GIY	215	СТУ	изр	PIU	СТУ	22.0	GIĀ	Gry	Gly	GIY
	Dro		Val	Dro	Dro	Glu		Dro	Gln	LAu	Sar		Dho	Δrα	Lys	Ser
	225	GIY	VUL	110	110	230	Olu	110	OIII	цси	235	Cys	1 110	nr 9	БуЗ	240
		Len	Ser	Asn	Val		Cvs	Glu	Trp	Glv		Arσ	Ser	Thr	Pro	
322		Dea	001		245	, 41	010	0		250		5	001		255	
	Leu	Thr	Thr	Lvs		Val	Leu	Leu	Val			Phe	Gln	Asn	Ser	Pro
326				260					265	5	-1-			270		
	Ala	Glu	Asp		Gln	Glu	Pro	Cys		Tyr	Ser	Gln	Glu	Ser	Gln	Lys
330													285			-1-
333	Phe	Ser	Cys	Gln	Leu	Ala	Val	Pro	Glu	Gly	Asp	Ser	Ser	Phe	Tyr	Ile
334		290	•				295			_	-	300			•	
337	Val		Met	Cys	Val	Ala	Ser	Ser	Val	Gly	Ser	Lys	Phe	Ser	Lys	Thr
	305			_		310				_	315	_			_	320
341	Gln	Thr	Phe	Gln	Gly	Cys	Gly	Ile	Leu	Gln	Pro	Asp	Pro	Pro	Ala	Asn
342					325					330					335	
345	Ile	Thr	Val	Thr	Ala	Val	Ala	Arg	Asn	Pro	Arg	Trp	Leu	Ser	Val	Thr
346				340					345					350		
349	${\tt Trp}$	Gln	Asp	${\tt Pro}$	His	Ser	Trp	Asn	Ser	Ser	Phe	Tyr	Arg	Leu	Arg	Phe

RAW SEQUENCE LISTING DATE: 07/30/2001

PATENT APPLICATION: US/09/462,416 TIME: 14:57:53

Input Set : A:\PTO_VSK.txt
Output Set: N:\CRF3\07302001\I462416.raw

350	355 360 365												
	Glu Leu Arg Tyr Arg Ala Glu Arg Ser Lys Thr Phe Thr Thr Trp Met												
354													
	Val Lys Asp Leu Gln His His Cys Val Ile His Asp Ala Trp Ser Gly 385 390 395 400												
	Leu Arg His Val Val Gln Leu Arg Ala Gln Glu Phe Gly Gln Gly												
362	405 410 415												
	Glu Trp Ser Glu Trp Ser Pro Glu Ala Met Gly Thr Pro Trp Thr Glu												
366	420 425 430												
	Ser Arg Ser Pro Pro Ala Glu Asn Glu Val Ser Thr Pro Met Gln Ala												
370	435 440 445												
	Leu Thr Thr Asn Lys Asp Asp Asp Ile Leu Phe Arg Asp Ser Ala												
374	450 455 460 Asn Ala Thr Ser Leu Pro Val												
	465 470												
	<210> SEQ ID NO: 9												
	<211> LENGTH: 24												
383	<212> TYPE: DNA												
	<213> ORGANISM: Artificial Sequence												
	<220> FEATURE:												
	<223> OTHER INFORMATION: synthetic												
	<400> SEQUENCE: 9	24											
	gcgacaagcc tcccagtgga attc	24											
	93 <210> SEQ ID NO: 10 94 <211> LENGTH: 18												
	<212> TYPE: DNA												
396	96 <213> ORGANISM: Artificial Sequence.												
	98 <220> FEATURE:												
	9 <223> OTHER INFORMATION: synthetic												
	<400> SEQUENCE: 10	1.0											
	cagtacccga attcatgc <210> SEQ ID NO: 11	18											
	<211> LENGTH: 31												
	<212> TYPE: DNA												
	<213> ORGANISM: Artificial Sequence												
	<220> FEATURE:												
411	<223> OTHER INFORMATION: synthetic												
	<400> SEQUENCE: 11												
	catggceegg geeeteetee eeegeegeee g	31											
	<210> SEQ ID NO: 12												
	<211> LENGTH: 22 <212> TYPE: DNA												
	<213> ORGANISM: Artificial Sequence												
	2 <220> FEATURE:												
	<223> OTHER INFORMATION: synthetic												
425	<400> SEQUENCE: 12												
426	gateceegee acceegggee ca	22											

VERIFICATION SUMMARY

ين 🐧 ۾ ايوان

PATENT APPLICATION: US/09/462,416

DATE: 07/30/2001

TIME: 14:57:54

Input Set : A:\PTO_VSK.txt
Output Set: N:\CRF3\07302001\I462416.raw